

SEQUENCE LISTING

<110> Millennium Pharmaceuticals, Inc. Curtis, Rory A.J.

<120> NOVEL MEMBERS OF THE CAPSAICIN/VANILLOID RECEPTOR FAMILY OF PROTEINS AND USES THEREOF

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geoggteagt caettagteg teagategee egteetggta teacagtget tetgtteagg 1080

| ttgcacactg ggcca | acagag gatccagcaa g | g atg aag aaa tgg Met Lys Lys Trp 1 | |
|------------------|---|---|---|
| | gct gcg gac cca ct Ala Ala Asp Pro Le 15 | | |
| | gac cct aac tcc ag Asp Pro Asn Ser Ar 30 | - | |
| | aag agc cgc acc cg Lys Ser Arg Thr Ar 45 | | |
| | ccg gtg gat tgc cc Pro Val Asp Cys Pr 60 | | _ |
| | atc aca gtc agc cc Ile Thr Val Ser Pr 8 | o Val Ile Thr Ile | |
| | acc ggt gcc agg ct Thr Gly Ala Arg Le 95 | | |
| | aag acc ctc agg ct Lys Thr Leu Arg Le 110 | | |
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| | agc aag aag cac ct Ser Lys Lys His Le 140 | | _ |
| | aag acc tgt ctg ct Lys Thr Cys Leu Le 16 | u Lys Ala Met Leu | |
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| | aag gag ctt gtc aa Lys Glu Leu Val As 190 | | |
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| ttc ggt Phe Gly | | | | | | | | | | | | | | | 1901 |
| atc gtg Ile Val 265 | _ | | - | _ | - | | | | _ | _ | - | - | | _ | 1949 |
| gcc agg Ala Arg 280 | _ | _ | | | | _ | | _ | | _ | | | | | 1997 |
| gcc gac Ala Asp | | - | - | _ | | _ | _ | | | _ | _ | - | | | 2045 |
| gag att Glu Ile | _ | _ | _ | | _ | | _ | | _ | _ | _ | | _ | | 2093 |
| gag ctc Glu Leu | | | | | | | | | | | | | | | 2141 |
| acc ggg Thr Gly 345 | | | | | | | | | | | | | | | 2189 |
| gag ccc Glu Pro 360 | | _ | | | _ | | | - | | | | | _ | | 2237 |
| ggg ccc Gly Pro | _ | | | _ | - | | | | | | | | | | 2285 |
| gag aag Glu Lys | | | | | | | | | | | | | | | 2333 |
| cct aat Pro Asn | - | | _ | _ | | - | | | - | _ | | _ | | _ | 2381 |
| cag gac Gln Asp 425 | _ | | _ | - | | | _ | _ | | | | | | | 2429 |
| ctg gtc Leu Val 440 | | _ | _ | | _ | | | | | _ | _ | _ | | | 2477 |
| agg ccc Arg Pro | | - | | _ | | | | _ | _ | _ | | | | _ | 2525 |
| tat ttc Tyr Phe | _ | _ | | | | | _ | | | | | | - | | 2573 |

| | | | _ | | | cag Gln | | | _ | _ | | | | | | 2621 | |
|---|---|---|---|---|---|-------------------|---|---|---|---|---|---|---|---|---|--------|--|
| | | | | | | agc Ser 510 | | | | | | | | | | 2669 | |
| | _ | | _ | _ | - | acc Thr | | | _ | | | _ | | | | 2717 | |
| | | | _ | | _ | gta Val | | | _ | - | _ | | | | | 2765 | |
| _ | | | | | | ggt Gly | | - | - | - | | | | | | 2813 . | |
| _ | | | _ | _ | | ctg Leu | _ | _ | _ | - | _ | | _ | | | 2861 | |
| | | _ | | _ | | ggg Gly 590 | | | | | | | _ | _ | | 2909 | |
| | | | | | | tcc Ser | | | | | | | | | | 2957 | |
| | | | | - | _ | agg Arg | | | _ | _ | | | | _ | _ | 3005 | |
| | | | - | _ | | ctg Leu | | _ | | | | | | | | 3053 | |
| - | _ | | | | | tat Tyr | | | | | | | | | | 3101 | |
| - | _ | _ | | _ | | ctc Leu 670 | | | | | _ | | | _ | | 3149 | |
| | _ | | _ | | | act Thr | - | | | | - | | | | | 3197 | |
| | | | | | | aga Arg | | | | | | | | | | 3245 | |
| _ | | | _ | _ | - | agg Arg | | | | | | | _ | _ | _ | 3293 | |
| | | | | | | gat Asp | | | | | | | | | | 3341 | |

730 735 740

| agg gtg gac gag gtg aac tgg acc acc tgg aac acc aac gtg ggc atc 3 Arg Val Asp Glu Val Asn Trp Thr Trp Asn Thr Asn Val Gly Ile 745 750 755 | 389 |
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| atc aac gaa gac ccg ggc aac tgt gag ggc gtc aag cgc acc ctg agc 3 Ile Asn Glu Asp Pro Gly Asn Cys Glu Gly Val Lys Arg Thr Leu Ser 760 765 770 775 | 437 |
| ttc tcc ctg cgg tca agc aga gtt tca ggc aga cac tgg aag aac ttt 3 Phe Ser Leu Arg Ser Ser Arg Val Ser Gly Arg His Trp Lys Asn Phe 780 785 790 | 485 |
| gcc ctg gtc ccc ctt tta aga gag gca agt gct cga gat agg cag tct 3 Ala Leu Val Pro Leu Leu Arg Glu Ala Ser Ala Arg Asp Arg Gln Ser 795 800 805 | 533 |
| gct cag ccc gag gaa gtt tat ctg cga cag ttt tca ggg tct ctg aag 3 Ala Gln Pro Glu Glu Val Tyr Leu Arg Gln Phe Ser Gly Ser Leu Lys 810 815 820 | 581 |
| cca gag gac gct gag gtc ttc aag agt cct gcc gct tcc ggg gag aag 3 Pro Glu Asp Ala Glu Val Phe Lys Ser Pro Ala Ala Ser Gly Glu Lys 825 830 835 | 629 |
| tgaggacgtc acgcagacag cactgtcaac actgggcctt aggagacccc gttgccacgg 3 ggggctgctg agggaacacc agtgctctgt cagcagcctg gcctggtctg tgcctgcca 3 gcatgttccc aaatctgtgc tggacaagct gtgggaagcg ttcttggaag catggggagt 3 gatgtacatc caaccgtcac tgtccccaag tgaatctcct aacagacttt caggttttta 3 ctcactttac taaaaaaaaa aaaaaaggg cggccgctta | 7 49 809 |
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Gly Ala Asp Val Gln Ala Ala Ala His Gly Asp Phe Phe Lys Lys Thr
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Lys Gly Arg Pro Gly Phe Tyr Phe Gly Glu Leu Pro Leu Ser Leu Ala
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Ala Cys Thr Asn Gln Leu Gly Ile Val Lys Phe Leu Leu Gln Asn Ser
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Trp Gln Thr Ala Asp Ile Ser Ala Arg Asp Ser Val Gly Asn Thr Val
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Leu His Ala Leu Val Glu Val Ala Asp Asn Thr Ala Asp Asn Thr Lys
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Phe Val Thr Ser Met Tyr Asn Glu Ile Leu Met Leu Gly Ala Lys Leu
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His Pro Thr Leu Lys Leu Glu Glu Leu Thr Asn Lys Lys Gly Met Thr
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Pro Leu Ala Leu Ala Gly Thr Gly Lys Ile Gly Val Leu Ala Tyr
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Ile Leu Gln Arg Glu Ile Gln Glu Pro Glu Cys Arg His Leu Ser Arg
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                                               365
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Ala Tyr Ser Ser Ser Glu Thr Pro Asn Arg His Asp Met Leu Leu Val
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                                   410
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Phe Thr Met Ala Ala Tyr Tyr Arg Pro Val Asp Gly Leu Pro Pro Phe
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Lys Met Glu Lys Ile Gly Asp Tyr Phe Arg Val Thr Gly Glu Ile Leu
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Ser Val Leu Gly Gly Val Tyr Phe Phe Phe Arg Gly Ile Gln Tyr Phe
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Glu Met Leu Phe Phe Leu Gln Ser Leu Phe Met Leu Ala Thr Val Val
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Leu Tyr Phe Ser His Leu Lys Glu Tyr Val Ala Ser Met Val Phe Ser
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Leu Ala Leu Gly Trp Thr Asn Met Leu Tyr Tyr Thr Arg Gly Phe Gln
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Gln Met Gly Ile Tyr Ala Val Met Ile Glu Lys Met Ile Leu Arg Asp
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Leu Cys Arg Phe Met Phe Val Tyr Ile Val Phe Leu Phe Gly Phe Ser
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                                585
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Thr Ala Val Val Thr Leu Ile Glu Asp Gly Lys Asn Asp Ser Leu Pro
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Ser Glu Ser Thr Ser His Arg Trp Arg Gly Pro Ala Cys Arg Pro Pro
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Asp Ser Ser Tyr Asn Ser Leu Tyr Ser Thr Cys Leu Glu Leu Phe Lys
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Phe Thr Ile Gly Met Gly Asp Leu Glu Phe Thr Glu Asn Tyr Asp Phe
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Lys Ala Val Phe Ile Ile Leu Leu Leu Ala Tyr Val Ile Leu Thr Tyr
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Lys Ile Ala Gln Glu Ser Lys Asn Ile Trp Lys Leu Gln Arg Ala Ile

| 11 | 5 | | 120 | | | | | 125 | | | | |
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| gat ctg ga Asp Leu Gl 130 | | _ | | | | | | | | | | 432 |
| aca gac aa Thr Asp As: 145 | | - | | | | | _ | | _ | - | - | 480 |
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| ctc ctg gag Leu Leu Gl | | | _ | - | | _ | _ | | | _ | | 576 |
| gcc agc tag Ala Ser Ty: 19 | r Thr Asp | - | | _ | | _ | | _ | _ | | | 624 |
| gcc atc gag Ala Ile Gli 210 | | - | | | | | | _ | | | | 672 |
| gga gca gad Gly Ala Asp 225 | | | _ | | | | | | | | | 720 |
| aaa ggg cgg Lys Gly Arg | | | | | - | - | | | | | | 768 |
| gcg tgc acc Ala Cys Th | _ | | | | | | | | | | | 816 |
| tgg cag acg Trp Gln Th 27! | r Ala Asp | _ | _ | | _ | _ | | | | _ | _ | 864 |
| ctg cac gcc Leu His Ala 290 | | | Ala | _ | | Thr | - | - | | _ | - | 912 |
| ttt gtg acg Phe Val Thi 305 | | | | | | | | | | | | 960 |
| cac ccg acg His Pro Th | | | | | | | | | | | | 1008 |
| ccg ctg gct Pro Leu Ala | | | | | | | | | | | | 1056 |
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| _ | | | | | _ | tac Tyr 375 | | - | | | | | | 1152 |
|---|---|---|---|---|---|-------------------|-------|---|---|---|---|---|---|------|
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| | | | | | | acc Thr | | | | | | | _ | 1248 |
| | _ | _ | | _ | | ctg Leu | - | _ | | - | _ | _ | | 1296 |
| | | | | | | ttc Phe | | | | | | | | 1344 |
| | | | | | | tac Tyr 455 | | | | | | | | 1392 |
| _ | _ | _ | | | | gac Asp | | _ | - | | | | | 1440 |
| | | | | | | tac Tyr | | | | | | | | 1488 |
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| | _ | | | | _ | cag Gln | _ | | | | | _ | _ | 1584 |
| | | | | | | aag Lys 535 | | | | | | | | 1632 |
| | | | | | | aac Asn | | | | | | | | 1680 |
| | | | | | | gtc Val | | | | | | | | 1728 |
| | | | | | | gtc Val | | | | | | | | 1776 |
| | | | | | | att Ile | | | | | | | | 1824 |
| | | | _ | _ | | agg Arg | | | | | | | | 1872 |

| | 610 | | | | | 615 | | | | | 620 | | | | |
|---|-----|---|------------|---|---|-----|---|---|---|---|-----|-------------------|---|---|------|
| | | | | | | | | | | | | ctg Leu | | | 1920 |
| | | | | | | | | | | | | tat Tyr | | | 1968 |
| | | | | | | | | | | | | ctc Leu 670 | | | 2016 |
| | | _ | | | _ | | | _ | | _ | | act Thr | - | | 2064 |
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| | | | | | | | | | | | | gat Asp | | | 2208 |
| - | - | | | | | | | | | | | tgg Trp 750 | | | 2256 |
| | | | | | | | | | | | | aac Asn | | | 2304 |
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| | - | | | - | | | - | _ | - | | | aga Arg | _ | _ | 2400 |
| | | | | | | | | | | | | tat Tyr | | | 2448 |
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gcc tgc ggc cgc ttc ttc cag aag ggc caa ggg act tgc ttt tat ttc

984

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| | | | | | | _ | _ | _ | _ | | _ | _ | tgg Trp | _ | | 1032 |
| - | | | | _ | | | | | - | | _ | _ | ctg Leu | _ | _ | 1080 |
| | | | | | | | - | _ | | _ | | | atg Met | | _ | 1128 |
| | | | | | | | | | | | _ | _ | tat Tyr 270 | _ | | 1176 |
| | | | | | - | _ | | _ | | | | _ | ctt Leu | | _ | 1224 |
| _ | | | | | | | | | | | | | gcc Ala | | | 1272 |
| | | | | | | | | | _ | _ | | | ttt Phe | | | 1320 |
| | | | | | | | | | | | | | Gly ggg | | | 1368 |
| | | | | | | | | | | | | | gag Glu 350 | | | 1416 |
| | | _ | | | | _ | | | _ | _ | _ | _ | cac His | _ | | 1464 |
| _ | _ | _ | _ | _ | | | _ | | | _ | _ | _ | gcg Ala | | | 1512 |
| - | _ | | | | _ | | | | | | _ | _ | aat Asn | _ | | 1560 |
| | _ | | | | | - | - | - | | | _ | | acc Thr | _ | _ | 1608 |
| | | _ | _ | | | _ | | | | - | | | tcc Ser 430 | _ | - | 1656 |
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| ggc c Gly G | | | | | | | | | | | | | | | | 1752 |
|-----------------------|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|------|
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| ccc c Pro L | _ | | | | | _ | _ | _ | | | | | | | | 1896 |
| tat a Tyr T | | | | | | | | | | | | | | | | 1944 |
| aag g Lys V 5 | | | | | | | | | | | | | | | | 1992 |
| ttc c Phe L 545 | | | | | - | | - | | | | | | | | | 2040 |
| tgg c | _ | | _ | _ | | | | | | _ | | | | | | 2088 |
| ccc a Pro M | _ | | | _ | | _ | - | | | | | | | | | 2136 |
| atc c | _ | _ | _ | | _ | | | | | | | | | | | 2184 |
| gag c Glu L 6 | _ | _ | | _ | | _ | _ | | | _ | | _ | | | | 2232 |
| ctg c Leu L 625 | | | | | | | | | | | | | | | | 2280 |
| ctc a Leu I | | _ | | _ | _ | | | _ | | _ | _ | _ | | - | | 2328 |
| tgg a Trp S | | | | | | | | _ | | | | | | | | 2376 |
| aat g Asn G | | | | | | | | | | | | | | | | 2424 |
| acc g | gtt | ggc | act | aag | cca | gat | ggc | agc | ccg | gat | gag | cgc | tgg | tgc | ttc | 2472 |

| Thr Val Gly Thr Lys Pro Asp Gly Ser Pro Asp Glu Arg Trp Cys Phe 690 695 700 |
|---|
| agg gtg gag gag gtg aac tgg gct tca tgg gag cag acg ctg cct acg Arg Val Glu Val Asn Trp Ala Ser Trp Glu Gln Thr Leu Pro Thr 705 710 715 720 |
| ctg tgt gag gac ccg tca ggg gca ggt gtc cct cga act ctc gag aac 2568 Leu Cys Glu Asp Pro Ser Gly Ala Gly Val Pro Arg Thr Leu Glu Asn 725 730 735 |
| cct gtc ctg gct tcc cct ccc aag gag gat gag gat ggt gcc tct gag Pro Val Leu Ala Ser Pro Pro Lys Glu Asp Glu Asp Gly Ala Ser Glu 740 745 750 |
| gaa aac tat gtg ccc gtc cag ctc ctc cag tcc aac tgatggccca 2662 Glu Asn Tyr Val Pro Val Gln Leu Leu Gln Ser Asn 755 760 |
| gatgcagcag gaggccagag gacagagcag aggatettte caaccacate tgetggetet 2722 ggggteecag tgaattetgg tggcaaatat atattteac taactcaaaa aaaaaaaaa 2782 aaaaaaaaaa aaaaaaaaa aaaaaaaa 2809 |
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| 1 5 10 15 Gly Gly Gln Glu Asp Gly Ser Glu Ala Asp Arg Gly Lys Leu Asp Phe |
| 20 25 30 . Gly Ser Gly Leu Pro Pro Met Glu Ser Gln Phe Gln Gly Glu Asp Arg 35 40 45 |
| Lys Phe Ala Pro Gln Ile Arg Val Asn Leu Asn Tyr Arg Lys Gly Thr 50 55 60 |
| Gly Ala Ser Gln Pro Asp Pro Asn Arg Phe Asp Arg Asp Arg Leu Phe 65 70 75 80 |
| Asn Ala Val Ser Arg Gly Val Pro Glu Asp Leu Ala Gly Leu Pro Glu 85 90 95 |
| Tyr Leu Ser Lys Thr Ser Lys Tyr Leu Thr Asp Ser Glu Tyr Thr Glu 100 105 110 |
| Gly Ser Thr Gly Lys Thr Cys Leu Met Lys Ala Val Leu Asn Leu Lys 115 120 125 |
| Asp Gly Val Asn Ala Cys Ile Leu Pro Leu Leu Gln Ile Asp Arg Asp 130 135 140 |
| Ser Gly Asn Pro Gln Pro Leu Val Asn Ala Gln Cys Thr Asp Asp Tyr 145 150 155 160 |
| Tyr Arg Gly His Ser Ala Leu His Ile Ala Ile Glu Lys Arg Ser Leu 165 170 175 |
| Gln Cys Val Lys Leu Leu Val Glu Asn Gly Ala Asn Val His Ala Arg |
| 180 185 190 |
| Ala Cys Gly Arg Phe Phe Gln Lys Gly Gln Gly Thr Cys Phe Tyr Phe 195 200 205 |
| Ala Cys Gly Arg Phe Phe Gln Lys Gly Gln Gly Thr Cys Phe Tyr Phe |
| Ala Cys Gly Arg Phe Phe Gln Lys Gly Gln Gly Thr Cys Phe Tyr Phe 195 200 205 Gly Glu Leu Pro Leu Ser Leu Ala Ala Cys Thr Lys Gln Trp Asp Val |
| Ala Cys Gly Arg Phe Phe Gln Lys Gly Gln Gly Thr Cys Phe Tyr Phe 195 Gly Glu Leu Pro Leu Ser Leu Ala Ala Cys Thr Lys Gln Trp Asp Val 210 Val Ser Tyr Leu Leu Glu Asn Pro His Gln Pro Ala Ser Leu Gln Ala |

f

```
260
                                265
Leu Leu Gln Ala Gly Ala Arg Leu Cys Pro Thr Val Gln Leu Glu Asp
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Ile Arg Asn Leu Gln Asp Leu Thr Pro Leu Lys Leu Ala Ala Lys Glu
                       295
                                            300
Gly Lys Ile Glu Ile Phe Arg His Ile Leu Gln Arg Glu Phe Ser Gly
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Leu Ser His Leu Ser Arg Lys Phe Thr Glu Trp Cys Tyr Gly Pro Val
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Arg Val Ser Leu Tyr Asp Leu Ala Ser Val Asp Ser Cys Glu Glu Asn
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Arg Met Val Val Leu Glu Pro Leu Asn Lys Leu Leu Gln Ala Lys Trp
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Asp Leu Leu Ile Pro Lys Phe Phe Leu Asn Phe Leu Cys Asn Leu Ile
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Tyr Met Phe Ile Phe Thr Ala Val Ala Tyr His Gln Pro Thr Leu Lys
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                                    410
Lys Gln Ala Ala Pro His Leu Lys Ala Glu Val Gly Asn Ser Met Leu
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                                425
                                                    430
Leu Thr Gly His Ile Leu Ile Leu Leu Gly Gly Ile Tyr Leu Leu Val
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Gly Gln Leu Trp Tyr Phe Trp Arg Arg His Val Phe Ile Trp Ile Ser
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                                            460
Phe Ile Asp Ser Tyr Phe Glu Ile Leu Phe Leu Phe Gln Ala Leu Leu
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                                        475
Thr Val Val Ser Gln Val Leu Cys Phe Leu Ala Ile Glu Trp Tyr Leu
                485
                                    490
Pro Leu Leu Val Ser Ala Leu Val Leu Gly Trp Leu Asn Leu Leu Tyr
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Tyr Thr Arg Gly Phe Gln His Thr Gly Ile Tyr Ser Val Met Ile Gln
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Lys Val Ile Leu Arg Asp Leu Leu Arg Phe Leu Leu Ile Tyr Leu Val
                       535
                                           540
Phe Leu Phe Gly Phe Ala Val Ala Leu Val Ser Leu Ser Gln Glu Ala
                    550
                                        555
Trp Arg Pro Glu Ala Pro Thr Gly Pro Asn Ala Thr Glu Ser Val Gln
                565
                                    570
Pro Met Glu Gly Gln Glu Asp Glu Gly Asn Gly Ala Gln Tyr Arg Gly
                                585
Ile Leu Glu Ala Ser Leu Glu Leu Phe Lys Phe Thr Ile Gly Met Gly
                            600
                                                605
Glu Leu Ala Phe Gln Glu Gln Leu His Phe Arg Gly Met Val Leu Leu
                       615
                                            620
Leu Leu Leu Ala Tyr Val Leu Leu Thr Tyr Ile Leu Leu Leu Asn Met
                    630
                                        635
Leu Ile Ala Leu Met Ser Glu Thr Val Asn Ser Val Ala Thr Asp Ser
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                                    650
Trp Ser Ile Trp Lys Leu Gln Lys Ala Ile Ser Val Leu Glu Met Glu
           660
                               665
Asn Gly Tyr Trp Trp Cys Arg Lys Lys Gln Arg Ala Gly Val Met Leu
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                                               685
Thr Val Gly Thr Lys Pro Asp Gly Ser Pro Asp Glu Arg Trp Cys Phe
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                        695
Arg Val Glu Glu Val Asn Trp Ala Ser Trp Glu Gln Thr Leu Pro Thr
                                        715
Leu Cys Glu Asp Pro Ser Gly Ala Gly Val Pro Arg Thr Leu Glu Asn
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                                    730
Pro Val Leu Ala Ser Pro Pro Lys Glu Asp Glu Asp Gly Ala Ser Glu
                                745
Glu Asn Tyr Val Pro Val Gln Leu Leu Gln Ser Asn
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755 760

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|------------|--------------------------------|-----|--------|-----|-----|-------------------|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| | 1> CI | | . (229 | 92) | | | | | | | | | | | | |
| atg | | | | | _ | tct Ser | | _ | | | _ | | | | _ | 48 |
| | | | | | | tct Ser | | | | | | | | | | 96 |
| | | | | | | atg Met | | | - | | | | | - | | 144 |
| | | _ | | _ | | aga Arg 55 | _ | | | | | _ | _ | | | 192 |
| | | | _ | _ | _ | cca Pro | | _ | | _ | - | _ | | | | 240 |
| | | | | | | gtc Val | | | _ | _ | _ | | | | | 288 |
| | | | _ | | _ | aag Lys | | | | _ | | _ | | | | 336 |
| _ | | | | | | tgc Cys | | | | | | | | | | 384 |
| | _ | _ | | | | att Ile 135 | | | | | | | | | | 432 |
| | | | | _ | | ctg Leu | _ | | _ | _ | _ | | _ | - | | 480 |
| | | | | | | ctg Leu | | | | | | _ | | _ | | 528 |
| | | | | | | gtg Val | | | | | | | | | | 576 |
| gcc | tgc | ggc | cgc | ttc | ttc | cag | aag | ggc | caa | ggg | act | tgc | ttt | tat | ttc | 624 |

| Ala | Cys | Gly 195 | Arg | Phe | Phe | Gln | Lys 200 | Gly | Gln | Gly | Thr | Cys 205 | Phe | Tyr | Phe | |
|-----|-----|------------|-----|-----|-----|-------------------|------------|-----|-----|-----|-----|------------|-----|-----|-----|------|
| | | | | | | ttg Leu 215 | | | | | | | | | | 672 |
| - | _ | | | - | | aac Asn | | | _ | | _ | | | | | 720 |
| | _ | | _ | | | aca Thr | - | _ | | | | | | | | 768 |
| _ | | | - | | | att Ile | - | - | | | _ | - | | _ | | 816 |
| | | | _ | | _ | cgc Arg | | - | | | | | | | | 864 |
| | | | | | | ctc Leu 295 | | | | | | | | | | 912 |
| | _ | | | | | agg Arg | | | _ | _ | | - | | | | 960 |
| _ | _ | | | | - | aag Lys | | | | | | | | | | 1008 |
| | | | | | | ctg Leu | | | | | | | | | | 1056 |
| | | | | | | gcc Ala | | | | | | | | | | 1104 |
| _ | _ | _ | | _ | | ccc Pro 375 | | | | | | | | | | 1152 |
| | | | | | | ttc Phe | | | | | | | | | | 1200 |
| | _ | | | | | gct Ala | _ | _ | | | | | | | | 1248 |
| _ | _ | - | - | | | ctg Leu | | | | | | | | | | 1296 |
| _ | _ | | | | | atc Ile | | | | | | | | | _ | 1344 |

| | | | | | | tgg Trp 455 | | | | | | | | | | 1392 |
|-----|-----|-----|-----|-----|-----|-------------------|-----|-----|-----|-----|-----|-----|-----|-----|-----|------|
| | | | | | | gaa Glu | | | | | | | | | | 1440 |
| | | | | | | ctg Leu | | | | | | | | | | 1488 |
| | _ | | | | | ctg Leu | | | | | _ | | _ | | | 1536 |
| | | _ | | | _ | cac His | | | | | | | | | | 1584 |
| - | _ | | _ | | _ | ctg Leu 535 | _ | - | | | _ | | | | _ | 1632 |
| | | | | | ~ | gta Val | _ | _ | | - | _ | - | _ | | _ | 1680 |
| | | | | | | aca Thr | | | | | | | | | | 1728 |
| | _ | | | _ | | gac Asp | _ | | | | _ | _ | | | | 1776 |
| | _ | - | _ | | _ | gag Glu | | | | | | | _ | | _ | 1824 |
| | _ | - | | _ | _ | cag Gln 615 | _ | | | _ | | - | | | | 1872 |
| - | _ | _ | _ | | | ctg Leu | | | | | _ | - | | | _ | 1920 |
| | | | | | | gag Glu | | | | | | | | | | 1968 |
| | _ | | | _ | _ | cag Gln | | _ | | | ~ | _ | | _ | | 2016 |
| | | | | | | agg Arg | | | | | | | | | | 2064 |
| acc | gtt | ggc | act | aag | cca | gat | ggc | agc | ccg | gat | gag | cgc | tgg | tgc | ttc | 2112 |

| Thr Val Gly Thr Lys Pro Asp Gly Ser Pro Asp Glu Arg Trp Cys Phe 690 695 700 | |
|---|-------------------------|
| agg gtg gag gag gtg aac tgg gct tca tgg gag cag acg ctg cct acg Arg Val Glu Val Asn Trp Ala Ser Trp Glu Gln Thr Leu Pro Thr 705 710 715 720 | 2160 |
| ctg tgt gag gac ccg tca ggg gca ggt gtc cct cga act ctc gag aac Leu Cys Glu Asp Pro Ser Gly Ala Gly Val Pro Arg Thr Leu Glu Asn 725 730 735 | 2208 |
| cct gtc ctg gct tcc cct ccc aag gag gat gag gat ggt gcc tct gag Pro Val Leu Ala Ser Pro Pro Lys Glu Asp Glu Asp Gly Ala Ser Glu 740 745 750 | 2256 |
| gaa aac tat gtg ccc gtc cag ctc ctc cag tcc aac Glu Asn Tyr Val Pro Val Gln Leu Leu Gln Ser Asn 755 760 | 2292 |
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| Gly Arg Phe Phe Gln Lys Gly Gln Gly Thr Cys Phe Tyr Phe Gly | 47 95 |
| Gly Arg Phe Phe Gln Lys Gly Gln Gly Thr Cys Phe Tyr Phe Gly 1 5 10 15 gag cta ccc ctc tct ttg gcc gct tgc acc aag cag tgg gat gtg gta Glu Leu Pro Leu Ser Leu Ala Ala Cys Thr Lys Gln Trp Asp Val Val | |
| Gly Arg Phe Phe Gln Lys Gly Gln Gly Thr Cys Phe Tyr Phe Gly 1 5 10 15 gag cta ccc ctc tct ttg gcc gct tgc acc aag cag tgg gat gtg gta Glu Leu Pro Leu Ser Leu Ala Ala Cys Thr Lys Gln Trp Asp Val Val 20 25 30 agc tac ctc ctg gag aac cca cac cag ccc gcc agc ctg cag gcc act Ser Tyr Leu Leu Glu Asn Pro His Gln Pro Ala Ser Leu Gln Ala Thr | 95 |
| Gly Arg Phe Phe Gln Lys Gly Gln Gly Thr Cys Phe Tyr Phe Gly 1 5 10 15 gag cta ccc ctc tct ttg gcc gct tgc acc aag cag tgg gat gtg gta Glu Leu Pro Leu Ser Leu Ala Ala Cys Thr Lys Gln Trp Asp Val Val 20 25 30 agc tac ctc ctg gag aac cca cac cag ccc gcc agc ctg cag gcc act Ser Tyr Leu Leu Glu Asn Pro His Gln Pro Ala Ser Leu Gln Ala Thr 35 40 45 gac tcc cag ggc aac aca gtc ctg cat gcc cta gtg atg atc tcg gac Asp Ser Gln Gly Asn Thr Val Leu His Ala Leu Val Met Ile Ser Asp | 95 143 |
| Gly Arg Phe Phe Gln Lys Gly Gln Gly Thr Cys Phe Tyr Phe Gly 1 5 10 15 gag cta ccc ctc tct ttg gcc gct tgc acc aag cag tgg gat gtg gta Glu Leu Pro Leu Ser Leu Ala Ala Cys Thr Lys Gln Trp Asp Val Val 20 25 30 agc tac ctc ctg gag aac cca cac cag ccc gcc agc ctg cag gcc act Ser Tyr Leu Leu Glu Asn Pro His Gln Pro Ala Ser Leu Gln Ala Thr 35 40 45 gac tcc cag ggc aac aca gtc ctg cat gcc cta gtg atg atc tcg gac Asp Ser Gln Gly Asn Thr Val Leu His Ala Leu Val Met Ile Ser Asp 50 55 60 aac tca gct gag aac att gca ctg gtg acc agc atg tat gat ggg ctc Asn Ser Ala Glu Asn Ile Ala Leu Val Thr Ser Met Tyr Asp Gly Leu | 95 143 191 |
| Gly Arg Phe Phe Gln Lys Gly Gln Gly Thr Cys Phe Tyr Phe Gly 1 5 10 15 gag cta ccc ctc tct ttg gcc gct tgc acc aag cag tgg gat gtg gta Glu Leu Pro Leu Ser Leu Ala Ala Cys Thr Lys Gln Trp Asp Val Val 20 25 30 agc tac ctc ctg gag aac cca cac cag ccc gcc agc ctg cag gcc act Ser Tyr Leu Leu Glu Asn Pro His Gln Pro Ala Ser Leu Gln Ala Thr 35 40 45 gac tcc cag ggc aac aca gtc ctg cat gcc cta gtg atg atc tcg gac Asp Ser Gln Gly Asn Thr Val Leu His Ala Leu Val Met Ile Ser Asp 50 55 60 aac tca gct gag aac att gca ctg gtg acc agc atg tat gat ggg ctc Asn Ser Ala Glu Asn Ile Ala Leu Val Thr Ser Met Tyr Asp Gly Leu 65 70 75 ctc caa gct ggg gcc cgc ctc tgc cct acc gtg cag ctt gag gac atc Leu Gln Ala Gly Ala Arg Leu Cys Pro Thr Val Gln Leu Glu Asp Ile | 95 143 191 239 |

| - | | | | _ | aag Lys | | | | - | _ | | | | | | 431 |
|---|---|---|---|---|-------------------|---|---|---|---|---|---|---|---|---|---|-------------|
| | _ | | | | ctg Leu | | | | | | | | | | | 479 |
| | _ | | | | gcc Ala 165 | | | _ | _ | _ | _ | | | | | 527 |
| _ | - | _ | _ | | ccc Pro | | | | - | | - | | | | | 575 |
| _ | | | | _ | ttc Phe | | | | | - | _ | | _ | _ | | 623 |
| _ | | | | | gct Ala | _ | - | | | | | | _ | _ | _ | 67 1 |
| | | | | | ctg Leu | | | | | | | | | | | 719 |
| _ | | | | | atc Ile 245 | _ | | | | | | | | | | 767 |
| | | | | | tgg Ţrp | | | | | | | | _ | | _ | 815 |
| | _ | _ | | | gaa Glu | | | | _ | | _ | _ | | | | 863 |
| | | | _ | | ctg Leu | _ | | - | _ | | | | | _ | | 911 |
| | | | | | ctg Leu | | | | | | | | | | | 959 |
| | _ | | | _ | cac His 325 | | | | | | - | | | | | 1007 |
| | - | | | _ | ctg Leu | | | _ | | | | | | | | 1055 |
| | | | | | ggt Gly | | | | | | | | | | | 1103 |
| | | | | | cgc Arg | | | | | | | | | | | 1151 |

| 370 | 375 | | 380 | |
|---|------------------------------|--------------------|-------------------------------|------------|
| gct tca tgg gag c Ala Ser Trp Glu G 385 | | | | |
| gca ggt gtc cct c Ala Gly Val Pro A 400 | | _ | | |
| aag gag gat gag g Lys Glu Asp Glu A 4 | | | | _ |
| ctc ctc cag tcc a Leu Leu Gln Ser A 435 | | gatgcagcag ga | aggccagag gacaga | gcag 1350 |
| aggatettte caacea atatttteae taacte ageggaegeg tgggte | aaaa aaaaaaaaa | | | |
| <210> 8 <211> 436 <212> PRT <213> Homo Sapien | s | ` | | |
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| Gly Arg Phe Phe G | ln Lys Gly Gln 5 | Gly Thr Cys 10 | Phe Tyr Phe Gly | Glu |
| Leu Pro Leu Ser L 20 | eu Ala Ala Cys | Thr Lys Gln 25 | Trp Asp Val Val 30 | Ser |
| Tyr Leu Leu Glu A 35 | sn Pro His Gln 40 | Pro Ala Ser | Leu Gln Ala Thr 45 | Asp |
| Ser Gln Gly Asn T | | Ala Leu Val | 77 | Asn |
| Ser Ala Glu Asn I | le Ala Leu Val | Thr Ser Met 75 | Tyr Asp Gly Leu | Leu 80 |
| Gln Ala Gly Ala A | rg Leu Cys Pro | | Leu Glu Asp Ile | |
| Asn Leu Gln Asp L | eu Thr Pro Leu | Lys Leu Ala | Ala Lys Glu Gly 110 | Lys |
| Ile Glu Ile Phe A | rg His Ile Leu 120 | | | Ser |
| His Leu Ser Arg L | | Trp Cys Tyr | | Val |
| Ser Leu Tyr Asp L | eu Ala Ser Val | _ | | |
| 145 Leu Glu Ile Ile A | | _ | | 160 Met |
| Val Val Leu Glu P | 65 ro Leu Asn Lys | | | Leu |
| Leu Ile Pro Lys P | | 185 Phe Leu Cys | | Met |
| 195 Phe Ile Phe Thr A 210 | 200 la Val Ala Tyr 215 | His Gln Pro | 205 Thr Leu Lys Lys 220 | Gln |
| Ala Ala Pro His L | | Val Gly Asn | | Thr |
| 225 Gly His Ile Leu I | 230 le Leu Leu Gly | 235 Gly Ile Tyr | Leu Leu Val Gly | 240 Gln |
| | 45 | 250 | 255 | |
| 260 | | 265 | 270 | - |

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Asp Ser Tyr Phe Glu Ile Leu Phe Leu Phe Gln Ala Leu Leu Thr Val
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                             280
 Val Ser Gln Val Leu Cys Phe Leu Ala Ile Glu Trp Tyr Leu Pro Leu
                         295
                                             300
Leu Val Ser Ala Leu Val Leu Gly Trp Leu Asn Leu Leu Tyr Tyr Thr
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                                         315
Arg Gly Phe Gln His Thr Gly Ile Tyr Ser Val Met Ile Gln Lys Lys
                                     330
                 325
Ala Ile Ser Val Leu Glu Met Glu Asn Gly Tyr Trp Trp Cys Arg Lys
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                                 345
                                                     350
Lys Gln Arg Ala Gly Val Met Leu Thr Val Gly Thr Lys Pro Asp Gly
       355
                             360
                                                 365
Ser Pro Asp Glu Arg Trp Cys Phe Arg Val Glu Glu Val Asn Trp Ala
                        375
Ser Trp Glu Gln Thr Leu Pro Thr Leu Cys Glu Asp Pro Ser Gly Ala
                                         395
                    390
Gly Val Pro Arg Thr Leu Glu Asn Pro Val Leu Ala Ser Pro Pro Lys
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                                     410
Glu Asp Glu Asp Gly Ala Ser Glu Glu Asn Tyr Val Pro Val Gln Leu
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Leu Gln Ser Asn
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cta ccc ctc tct ttg gcc gct tgc acc aag cag tgg gat gtg gta agc
Leu Pro Leu Ser Leu Ala Ala Cys Thr Lys Gln Trp Asp Val Val Ser
             20
                                                                    144
tac ctc ctg gag aac cca cac cag ccc gcc agc ctg cag gcc act gac
Tyr Leu Leu Glu Asn Pro His Gln Pro Ala Ser Leu Gln Ala Thr Asp
         35
                              40
tcc cag ggc aac aca gtc ctg cat gcc cta gtg atg atc tcg gac aac
Ser Gln Gly Asn Thr Val Leu His Ala Leu Val Met Ile Ser Asp Asn
     50
                          55
                                              60
tca gct gag aac att gca ctg gtg acc agc atg tat gat ggg ctc ctc
                                                                    240
Ser Ala Glu Asn Ile Ala Leu Val Thr Ser Met Tyr Asp Gly Leu Leu
 65
                                                                    288
caa gct ggg gcc cgc ctc tgc cct acc gtg cag ctt gag gac atc cgc
Gln Ala Gly Ala Arg Leu Cys Pro Thr Val Gln Leu Glu Asp Ile Arg
aac ctg cag gat ctc acg cct ctg aag ctg gcc gcc aag gag ggc aag
Asn Leu Gln Asp Leu Thr Pro Leu Lys Leu Ala Ala Lys Glu Gly Lys
            100
                                 105
```

| | | | | | | atc Ile | | | | | | | _ | | | 384 |
|---|---|---|---|---|---|-------------------|---|---|---|---|---|---|---|---|---|------|
| | | | _ | - | | acc Thr 135 | | | | | | | | | | 432 |
| _ | _ | | - | _ | _ | tct Ser | | _ | _ | _ | | | | | | 480 |
| _ | | | | _ | | cat His | _ | _ | _ | _ | | _ | | _ | _ | 528 |
| _ | _ | _ | | | _ | aac Asn | | _ | _ | _ | | | | | | 576 |
| | | | _ | | | tta Leu | | | _ | _ | | _ | _ | | _ | 624 |
| | | | | | | gcc Ala 215 | | | | | | | | | | 672 |
| | | | | | | gcg Ala | | | | | | | | | | 720 |
| | | | | | | cta Leu | | | | | | | | | | 768 |
| | | | | | | cgc Arg | | | | | | | | | | 816 |
| _ | _ | | | _ | | ctc Leu | | _ | | _ | _ | _ | | | | 864 |
| | | | | | | ttc Phe 295 | | | _ | _ | | | | | | 912 |
| | | | | | | ctg Leu | | | | | | | | | | 960 |
| | | | | | | ggc | | | | | | | | | | 1008 |
| | | | | | | atg Met | | | | | | | | | | 1056 |
| - | | | | | | atg Met | | | | | | | | | | 1104 |

355 360 365 1152 age eeg gat gag ege tgg tge tte agg gtg gag gtg aac tgg get Ser Pro Asp Glu Arg Trp Cys Phe Arg Val Glu Val Asn Trp Ala 375 370 tca tgg gag cag acg ctg cct acg ctg tgt gag gac ccg tca ggg gca 1200 Ser Trp Glu Gln Thr Leu Pro Thr Leu Cys Glu Asp Pro Ser Gly Ala 385 390 395 ggt gtc cct cga act ctc gag aac cct gtc ctg gct tcc cct ccc aag 1248 Gly Val Pro Arg Thr Leu Glu Asn Pro Val Leu Ala Ser Pro Pro Lys gag gat gag gat ggt gcc tct gag gaa aac tat gtg ccc gtc cag ctc 1296 Glu Asp Glu Asp Gly Ala Ser Glu Glu Asn Tyr Val Pro Val Gln Leu 425 ctc cag tcc aac 1308 Leu Gln Ser Asn 435 <210> 10 <211> 1794 <212> DNA <213> Rattus sp. <220> <221> CDS <222> (2)...(1663) <400> 10 g teg acc cae geg tee get ett tet etg get geg tge acc aag cag tgg 49 Ser Thr His Ala Ser Ala Leu Ser Leu Ala Ala Cys Thr Lys Gln Trp gat gtg gtg acc tac ctc ctg gag aac cca cac cag ccg gcc agc ctg 97 Asp Val Val Thr Tyr Leu Leu Glu Asn Pro His Gln Pro Ala Ser Leu 25 2.0 gag gcc acc gac tcc ctg ggc aac aca gtc ctg cat gct ctg gta atg Glu Ala Thr Asp Ser Leu Gly Asn Thr Val Leu His Ala Leu Val Met 40 att gca gat aac tcg cct gag aac agt gcc ctg gtg atc cac atg tac Ile Ala Asp Asn Ser Pro Glu Asn Ser Ala Leu Val Ile His Met Tyr 55 gac ggg ctt cta caa atg ggg gcg cgc ctc tgc ccc act gtg cag ctt 241 Asp Gly Leu Leu Gln Met Gly Ala Arg Leu Cys Pro Thr Val Gln Leu gag gaa atc tcc aac cac caa ggc ctc aca ccc ctg aaa cta gcc gcc 289 Glu Glu Ile Ser Asn His Gln Gly Leu Thr Pro Leu Lys Leu Ala Ala 85 aag gaa ggc aaa atc gag att ttc agg cac att ctg cag cgg gaa ttc Lys Glu Gly Lys Ile Glu Ile Phe Arg His Ile Leu Gln Arg Glu Phe 100 105 110

tca gga ccg tac cag ccc ctt tcc cga aag ttt act gag tgg tgt tac

| Ser | Gly | Pro 115 | Tyr | Gln | Pro | Leu | Ser 120 | Arg | Lys | Phe | Thr | Glu 125 | Trp | Cys | Tyr | |
|-----|-----|------------|-----|-----|-----|-------------------|------------|-----|-----|-----|-----|------------|-----|-----|-----|------|
| | | - | | _ | _ | ctg Leu 135 | | _ | _ | | | | - | - | | 433 |
| | | | | | | gag Glu | | | | | | | | | | 481 |
| | | | - | | | gtt Val | | _ | | - | | | | | _ | 529 |
| | | | _ | | | gtc Val | | _ | | | | | | - | | 577 |
| | _ | - | | _ | | atc Ile | | | - | - | - | | | _ | | 625 |
| | _ | ~ | ~ | | _ | atc Ile 215 | | | | | | | | | _ | 673 |
| | _ | _ | - | - | | cac His | | | | | | | | | | 721 |
| | | _ | | _ | _ | tgg Trp | | | | | | _ | _ | | | 769 |
| | | | | - | | agc Ser | | | | | | | | | | 817 |
| _ | _ | | | | _ | tcc Ser | _ | | _ | _ | | _ | | | | 865 |
| ~ ~ | | | | _ | | gtg Val 295 | | | | | _ | | | _ | | 913 |
| | | | | | | ggc Gly | | | | | | | | | | 961 |
| _ | | _ | _ | _ | | ctt Leu | _ | _ | | | | | | | | 1009 |
| | _ | _ | | | | ggc Gly | | _ | _ | _ | | _ | _ | _ | - | 1057 |
| - | | | - | _ | | aaa Lys | _ | | - | | | | | | | 1105 |

| acg gaa cag ccc acg gtg ggc cag gag gag gag cca gct cca tat cgg 1 Thr Glu Gln Pro Thr Val Gly Gln Glu Glu Pro Ala Pro Tyr Arg 370 375 380 | 1153 |
|---|--------------|
| agc att ctg gat gcc tcc cta gag ctg ttc aag ttc acc att ggt atg Ser Ile Leu Asp Ala Ser Leu Glu Leu Phe Lys Phe Thr Ile Gly Met 385 390 395 400 | 1201 |
| ggg gag ctg gct ttc cag gaa cag ctg cgt ttt cgt ggg gtg gtc ctg 1 Gly Glu Leu Ala Phe Gln Glu Gln Leu Arg Phe Arg Gly Val Val Leu 405 410 415 | L249 |
| ctg ttg ctg ttg gcc tac gtc ctt ctc acc tac gtc ctg ctg ctc aac 1 Leu Leu Leu Leu Ala Tyr Val Leu Leu Thr Tyr Val Leu Leu Asn 420 430 | L29 7 |
| atg ctc att gct ctc atg agc gaa act gtc aac cac gtt gct gac aac 1 Met Leu Ile Ala Leu Met Ser Glu Thr Val Asn His Val Ala Asp Asn 435 440 445 | 1345 |
| agc tgg agc atc tgg aag ttg cag aaa gcc atc tct gtc ttg gag atg 1 Ser Trp Ser Ile Trp Lys Leu Gln Lys Ala Ile Ser Val Leu Glu Met 450 455 460 | L393 |
| gag aat ggt tac tgg tgc tgc cgg agg aag aaa cat cgt gaa ggg agg 1 Glu Asn Gly Tyr Trp Trp Cys Arg Arg Lys Lys His Arg Glu Gly Arg 465 470 475 480 | 1441 |
| ctg ctg aaa gtc ggc acc agg ggg gat ggt acc cct gat gag cgc tgg 1 Leu Leu Lys Val Gly Thr Arg Gly Asp Gly Thr Pro Asp Glu Arg Trp 485 490 495 | L489 |
| tgc ttc agg gtg gag gaa gta aat tgg gct gct tgg gag aag act ctt 1 Cys Phe Arg Val Glu Glu Val Asn Trp Ala Ala Trp Glu Lys Thr Leu 500 505 510 | L537 |
| ccc acc tta tct gag gat cca tca ggg cca ggc atc act ggt aat aaa 1 Pro Thr Leu Ser Glu Asp Pro Ser Gly Pro Gly Ile Thr Gly Asn Lys 515 520 525 | L585 |
| aag aac cca acc tct aaa ccg ggg aag aac agt gcc tca gag gaa gac 1 Lys Asn Pro Thr Ser Lys Pro Gly Lys Asn Ser Ala Ser Glu Glu Asp 530 535 540 | L633 |
| cat ctg ccc ctt cag gtc ctc cag tcc ccc tgatggccca gatgcagcag 1 His Leu Pro Leu Gln Val Leu Gln Ser Pro 545 550 | L683 |
| caggctggca ggatggagta gggaatcttc ccagccacac cagaggctac tgaattttgg 1 tggaaatata aatattttt ttgcataaaa aaaaaaaaa agggcggccg c 1 | L743 L794 |
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| 1 5 10 15 Asp Val Val Thr Tyr Leu Leu Glu Asn Pro His Gln Pro Ala Ser Leu 20 25 30 | |

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Ile Ala Asp Asn Ser Pro Glu Asn Ser Ala Leu Val Ile His Met Tyr
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                                            60
Asp Gly Leu Leu Gln Met Gly Ala Arg Leu Cys Pro Thr Val Gln Leu
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                   70
Glu Glu Ile Ser Asn His Gln Gly Leu Thr Pro Leu Lys Leu Ala Ala
                                    90
Lys Glu Gly Lys Ile Glu Ile Phe Arg His Ile Leu Gln Arg Glu Phe
           100
                               105
                                                   110
Ser Gly Pro Tyr Gln Pro Leu Ser Arg Lys Phe Thr Glu Trp Cys Tyr
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                           120
                                               125
Gly Pro Val Arg Val Ser Leu Tyr Asp Leu Ser Ser Val Asp Ser Trp
                       135
Glu Lys Asn Ser Val Leu Glu Ile Ile Ala Phe His Cys Lys Ser Pro
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                   150
Asn Arg His Arg Met Val Val Leu Glu Pro Leu Asn Lys Leu Leu Gln
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Glu Lys Trp Asp Arg Leu Val Ser Arg Phe Phe Phe Asn Phe Ala Cys
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Tyr Leu Val Tyr Met Phe Ile Phe Thr Val Val Ala Tyr His Gln Pro
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Ser Leu Asp Gln Pro Ala Ile Pro Ser Ser Lys Ala Thr Phe Gly Glu
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                                           220
Ser Met Leu Leu Gly His Ile Leu Ile Leu Gly Gly Ile Tyr
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                   230
Leu Leu Gly Gln Leu Trp Tyr Phe Trp Arg Arg Arg Leu Phe Ile
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                                   250
Trp Ile Ser Phe Met Asp Ser Tyr Phe Glu Ile Leu Phe Leu Leu Gln
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                               265
                                                    270
Ala Leu Leu Thr Val Leu Ser Gln Val Leu Arg Phe Met Glu Thr Glu
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Trp Tyr Leu Pro Leu Leu Val Leu Ser Leu Val Leu Gly Trp Leu Asn
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                                           300
Leu Leu Tyr Tyr Thr Arg Gly Phe Gln His Thr Gly Ile Tyr Ser Val
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Met Ile Gln Lys Val Ile Leu Arg Asp Leu Leu Arg Phe Leu Leu Val
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Tyr Leu Val Phe Leu Phe Gly Phe Ala Val Ala Leu Val Ser Leu Ser
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Arg Glu Ala Arg Ser Pro Lys Ala Pro Glu Asp Asn Asn Ser Thr Val
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Thr Glu Gln Pro Thr Val Gly Gln Glu Glu Pro Ala Pro Tyr Arg
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Ser Ile Leu Asp Ala Ser Leu Glu Leu Phe Lys Phe Thr Ile Gly Met
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Gly Glu Leu Ala Phe Gln Glu Gln Leu Arg Phe Arg Gly Val Val Leu
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Met Leu Ile Ala Leu Met Ser Glu Thr Val Asn His Val Ala Asp Asn
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Ser Trp Ser Ile Trp Lys Leu Gln Lys Ala Ile Ser Val Leu Glu Met
                       455
                                           460
Glu Asn Gly Tyr Trp Trp Cys Arg Lys Lys His Arg Glu Gly Arg
                   470
                                      475
Leu Leu Lys Val Gly Thr Arg Gly Asp Gly Thr Pro Asp Glu Arg Trp
                                   490
                485
Cys Phe Arg Val Glu Glu Val Asn Trp Ala Ala Trp Glu Lys Thr Leu
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           500
                                                    510
Pro Thr Leu Ser Glu Asp Pro Ser Gly Pro Gly Ile Thr Gly Asn Lys
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| | 180 | 185 | 5 | 190 | |
|---|-----|-----|---------------------------------------|-----|---|
| | _ | | gtc gtt gcc Val Val Ala | | |
| | | | a tca aaa gcg Ser Lys Ala 220 | | - |
| | | | g atc ctg ctt 1 Ile Leu Leu 235 | | |
| _ | | | tgg cgg cgg Trp Arg Arg 250 | | |
| | | - | gaa atc ctc Glu Ile Leu | | |
| | | | g ctg cgc ttc Leu Arg Phe | | |
| | - | | c cta gtg ctg Leu Val Leu 300 | | |
| | | | g cac aca ggc n His Thr Gly 315 | | |
| - | | | c ctg ctc cgt Leu Leu Arg 330 | | _ |
| | | | gta gcc cta a Val Ala Leu 5 | | |
| | | _ | gaa gat aac Glu Asp Asn | | |
| | | | g gag gag cca n Glu Glu Pro 380 | | |
| | - | | g ttc aag ttc 1 Phe Lys Phe 395 | | |
| | | | g cgt ttt cgt n Arg Phe Arg 410 | | |
| | | | e acc tac gtc I Thr Tyr Val | | |

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atg ctc att gct ctc atg agc gaa act gtc aac cac gtt gct gac aac
Met Leu Ile Ala Leu Met Ser Glu Thr Val Asn His Val Ala Asp Asn
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                            440
                                                                   1392
age tgg age ate tgg aag ttg cag aaa gee ate tet gte ttg gag atg
Ser Trp Ser Ile Trp Lys Leu Gln Lys Ala Ile Ser Val Leu Glu Met
    450
                        455
gag aat ggt tac tgg tgg tgc cgg agg aag aaa cat cgt gaa ggg agg
                                                                   1440
Glu Asn Gly Tyr Trp Trp Cys Arg Arg Lys Lys His Arg Glu Gly Arg
                    470
                                        475
465
ctg ctg aaa gtc ggc acc agg ggg gat ggt acc cct gat gag cgc tgg
                                                                   1488
Leu Leu Lys Val Gly Thr Arg Gly Asp Gly Thr Pro Asp Glu Arg Trp
                485
                                    490
tgc ttc agg gtg gag gaa gta aat tgg gct gct tgg gag aag act ctt
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Cys Phe Arg Val Glu Glu Val Asn Trp Ala Ala Trp Glu Lys Thr Leu
            500
                                505
                                                     510
                                                                   1584
ccc acc tta tct gag gat cca tca ggg cca ggc atc act ggt aat aaa
Pro Thr Leu Ser Glu Asp Pro Ser Gly Pro Gly Ile Thr Gly Asn Lys
                            520
                                                 525
                                                                   1632
aag aac cca acc tct aaa ccg ggg aag aac agt gcc tca gag gaa gac
Lys Asn Pro Thr Ser Lys Pro Gly Lys Asn Ser Ala Ser Glu Glu Asp
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                                                                   1662
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His Leu Pro Leu Gln Val Leu Gln Ser Pro
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Gly Ser Gly Leu Pro Pro Met Glu Ser Gln Phe Gln Gly Glu Asp Arg
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Lys Phe Ala Pro Gln Ile Arg Val Asn Leu Asn Tyr Arg Lys Gly Thr
                                            60
                        5.5
Gly Ala Ser Gln Pro Asp Pro Asn Arg Phe Asp Arg Asp Arg Leu Phe
                                       75
Asn Ala Val Ser Arg Gly Val Pro Glu Asp Leu Ala Gly Leu Pro Glu
               85
                                   90
Tyr Leu Ser Lys Thr Ser Lys Tyr Leu Thr Asp Ser Glu Tyr Thr Glu
                               105
           100
Gly Ser Thr Gly Lys Thr Cys Leu Met Lys Ala Val Leu Asn Leu Lys
                            120
Asp Gly Val Asn Ala Cys Ile Leu Pro Leu Leu Gln Ile Asp Arg Asp
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                                            140
   130
Ser Gly Asn Pro Gln Pro Leu Val Asn Ala Gln Cys Thr Asp Asp Tyr
                   150
                                        155
Tyr Arg Gly His Ser Ala Leu His Ile Ala Ile Glu Lys Arg Ser Leu
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                                   170
Gln Cys Val Lys Leu Leu Val Glu Asn Gly Ala Asn Val His Ala Arg
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                                                    190
           180
Ala Cys Gly Arg Phe Phe Gln Lys Gly Gln Gly Thr Cys Phe Tyr Phe
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                                                205
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Gly Glu Leu Pro Leu Ser Leu Ala Ala Cys Thr Lys Gln Trp Asp Val
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                        215
                                           220
Val Ser Tyr Leu Leu Glu Asn Pro His Gln Pro Ala Ser Leu Gln Ala
                                       235
                  230
Thr Asp Ser Gln Gly Asn Thr Val Leu His Ala Leu Val Met Ile Ser
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Asp Asn Ser Ala Glu Asn Ile Ala Leu Val Thr Ser Met Tyr Asp Gly
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Leu Leu Gln Ala Gly Ala Arg Leu Cys Pro Thr Val Gln Leu Glu Asp
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                                                285
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Ile Arg Asn Leu Gln Asp Leu Thr Pro Leu Lys Leu Ala Ala Lys Glu
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Gly Lys Ile Glu Ile Phe Arg His Ile Leu Gln Arg Glu Phe Ser Gly
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Leu Ser His Leu Ser Arg Lys Phe Thr Glu Trp Cys Tyr Gly Pro Val
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                                    330
Arg Val Ser Leu Tyr Asp Leu Ala Ser Val Asp Ser Cys Glu Glu Asn
                                345
Ser Val Leu Glu Ile Ile Ala Phe His Cys Lys Ser Pro His Arg His
       355
                            360
                                                365
Arg Met Val Val Leu Glu Pro Leu Asn Lys Leu Leu Gln Ala Lys Trp
                                            380.
                        375
Asp Leu Leu Ile Pro Lys Phe Phe Leu Asn Phe Leu Cys Asn Leu Ile
                                       395
                   390
Tyr Met Phe Ile Phe Thr Ala Val Ala Tyr His Gln Pro Thr Leu Lys
               405
                                   410
Lys Gln Ala Ala Pro His Leu Lys Ala Glu Val Gly Asn Ser Met Leu
                                425
Leu Thr Gly His Ile Leu Ile Leu Leu Gly Gly Ile Tyr Leu Leu Val
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                            440
                                                445
Gly Gln Leu Trp Tyr Phe Trp Arg Arg His Val Phe Ile Trp Ile Ser
                        455
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Phe Ile Asp Ser Tyr Phe Glu Ile Leu Phe Leu Phe Gln Ala Leu Leu
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Tyr Thr Arg Gly Phe Gln His Thr Gly Ile Tyr Ser Val Met Ile Gln
       515
                            520
                                                525
Lys Lys Ala Ile Ser Val Leu Glu Met Glu Asn Gly Tyr Trp Trp Cys
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Arg Lys Lys Gln Arg Ala Gly Val Met Leu Thr Val Gly Thr Lys Pro
                    550
                                        555
Asp Gly Ser Pro Asp Glu Arg Trp Cys Phe Arg Val Glu Glu Val Asn
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                565
Trp Ala Ser Trp Glu Gln Thr Leu Pro Thr Leu Cys Glu Asp Pro Ser
            580
                                585
                                                    590
Gly Ala Gly Val Pro Arg Thr Leu Glu Asn Pro Val Leu Ala Ser Pro
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                                                605
                            600
Pro Lys Glu Asp Glu Asp Gly Ala Ser Glu Glu Asn Tyr Val Pro Val
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Gln Leu Leu Gln Ser Asn
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                       Met Glu Gln Arg Ala Ser Leu Asp Ser Glu Glu
tot gag toe coa coe caa gag aac toe tge etg gac cot coa gac aga
                                                                   161
Ser Glu Ser Pro Pro Gln Glu Asn Ser Cys Leu Asp Pro Pro Asp Arg
                                 20
            ·15
gac cet aac tgc aag cca cet cca gtc aag ccc cac atc ttc act acc
                                                                   209
Asp Pro Asn Cys Lys Pro Pro Pro Val Lys Pro His Ile Phe Thr Thr
                             35
agg agt cgt acc cgg ctt ttt ggg aag ggt gac tcg gag gag gcc tct
Arg Ser Arg Thr Arg Leu Phe Gly Lys Gly Asp Ser Glu Glu Ala Ser
                         50
                                             55
ccc ctg gac tgc cct tat gag gaa ggc ggg ctg gct tcc tgc cct atc
Pro Leu Asp Cys Pro Tyr Glu Glu Gly Gly Leu Ala Ser Cys Pro Ile
                                                                   353
atc act gtc agc tct gtt cta act atc cag agg cct ggg gat gga cct
Ile Thr Val Ser Ser Val Leu Thr Ile Gln Arg Pro Gly Asp Gly Pro
gcc agt gtc agg ccg tca tcc cag gac tcc gtc tcc gct ggt gag aag
                                                                   401
Ala Ser Val Arg Pro Ser Ser Gln Asp Ser Val Ser Ala Gly Glu Lys
             95
                                100
ccc ccg agg ctc tat gat cgc agg agc atc ttc gat gct gtg gct cag
                                                                   449
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Thr Val Val Ser Gln Val Leu Cys Phe Leu Ala Ile Glu Trp Tyr Leu

| Pro | Pro | Arg 110 | Leu | Tyr | Asp | Arg | Arg 115 | Ser | Ile | Phe | Asp | Ala 120 | Val | Ala | Gln | |
|-----|-----|------------|-----|-----|-----|-----|-------------------|-----|-----|-----|-----|------------|-----|-----|-----|------|
| _ | | _ | _ | | _ | | agc Ser | _ | _ | | | _ | _ | | _ | 497 |
| _ | _ | _ | _ | | - | _ | gag Glu | | | _ | | | | | _ | 545 |
| | - | _ | | | _ | _ | ctc Leu | | _ | | | | | | | 593 |
| | | _ | _ | | _ | - | gtt Val | - | | _ | | _ | _ | _ | _ | 641 |
| ~ | | ~ | | _ | _ | | aca Thr 195 | _ | _ | | | - | | _ | | 689 |
| _ | _ | | | - | | - | cgg Arg | | | | | | | | | 737 |
| _ | | | | | - | - | gtc Val | - | _ | _ | | | | | | 785 |
| | | | | | | | cct Pro | | | | | | | | | 833 |
| _ | | | _ | | _ | | aac Asn | - | _ | - | | | | | | 881 |
| _ | _ | | | | _ | | gca Ala 275 | _ | | | _ | | _ | | | 929 |
| | | _ | | | | - | ctg Leu | | | | - | - | | | _ | 977 |
| | | | | | | | agc Ser | | | | | | | | | 1025 |
| | - | | | | | _ | ctg Leu | _ | _ | - | | | | | | 1073 |
| _ | | | _ | | _ | _ | ctg Leu | _ | _ | _ | _ | | _ | | | 1121 |
| _ | _ | - | | | | _ | agg Arg 355 | _ | | | | | - | | _ | 1169 |

| | c cta s Leu 365 | | | _ | | | _ | | - | | | | | | | 1217 |
|----|-----------------------|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|------|
| | c ctt r Leu 0 | | | | | | _ | | | | | | | | _ | 1265 |
| | g gag u Glu | | | - | | _ | _ | _ | - | | | | | | | 1313 |
| | g ctt t Leu | | | | | | | | | | | | | | | 1361 |
| _ | a ttt g Phe | - | _ | _ | | | | | | | | - | | - | _ | 1409 |
| | t atg r Met 445 | | | | | | - | _ | | | | | | | | 1457 |
| | g ccc u Pro O | | | | | | | | | | | | | | _ | 1505 |
| | c gga r Gly | | | _ | | | | | | - | | | | _ | - | 1553 |
| | g att y Ile | | | | _ | _ | | - | | | | | | | | 1601 |
| _ | g gac l Asp | _ | | _ | | | | | | _ | _ | _ | | | | 1649 |
| | g gtg u Val 525 | | | _ | _ | | | _ | | _ | | - | | | | 1697 |
| | c atg r Met D | | | | | | | | | | | | | | | 1745 |
| | c cga r Arg | | | _ | _ | _ | | | | _ | _ | _ | | | _ | 1793 |
| | g atc t Ile | | _ | _ | - | _ | | | _ | | _ | | | | | 1841 |
| | g ttt u Phe | | | | | | | | | | | | | | | 1889 |
| aa | t aac | tct | ctg | cct | atg | gag | tcc | aca | cca | cac | aag | tgc | cgg | ggg | tct | 1937 |

| Asn | Asn 605 | Ser | Leu | Pro | Met | Glu 610 | Ser | Thr | Pro | His | Lys 615 | Cys | Arg | Gly | Ser | |
|-----|------------|-------|------|------|------|-------------------|------|------|------|------|------------|------|-------------------|------|----------------------------|------|
| | | | | | | tct Ser | | | | | | | | | | 1985 |
| | | | | | | atc Ile | | | | | | | | | | 2033 |
| | | | | | | gtc Val | | | | | | | | | | 2081 |
| | | | | | | ctg Leu | | | | | | | | | | 2129 |
| ~ ~ | | _ | | _ | | gca Ala 690 | | | | _ | | | | _ | _ | 2177 |
| _ | _ | _ | | | | ctg Leu | _ | | | | | | | | | 2225 |
| | | | | | | tct Ser | | | | | | | | | | 2273 |
| | | | | | | tac Tyr | | | | | | | | | | 2321 |
| | | | | | | acc Thr | | _ | | | | | | | | 2369 |
| - | | | | | | aag Lys 770 | | | | | | | | | | 2417 |
| | | | | | | aac Asn | | | | | | | | | | 2465 |
| | | | | | | cga Arg | | | | | | | | | | 2513 |
| _ | | - | _ | | | acg Thr | | | | _ | | | | | | 2561 |
| - | | _ | - | | _ | gtc Val | | | - | | taa * | tgg | acac [.] | tat | | 2607 |
| tct | gggt | cca d | ctct | gtgc | ct g | cctag | ggca | c at | teet | agga | ctt | cggc | ggg (| cctg | acgtta ctgtgg tttggc | 2727 |

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<213> Rattus sp.
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                               25
Pro Pro Pro Val Lys Pro His Ile Phe Thr Thr Arg Ser Arg Thr Arg
                          40
                                               45
Leu Phe Gly Lys Gly Asp Ser Glu Glu Ala Ser Pro Leu Asp Cys Pro
                       55
Tyr Glu Glu Gly Gly Leu Ala Ser Cys Pro Ile Ile Thr Val Ser Ser
                    70
                                        75
Val Leu Thr Ile Gln Arg Pro Gly Asp Gly Pro Ala Ser Val Arg Pro
                                   90
               85
Ser Ser Gln Asp Ser Val Ser Ala Gly Glu Lys Pro Pro Arg Leu Tyr
                                105
                                                    110
Asp Arg Arg Ser Ile Phe Asp Ala Val Ala Gln Ser Asn Cys Gln Glu
       115
                           120
                                                125
Leu Glu Ser Leu Leu Pro Phe Leu Gln Arg Ser Lys Lys Arg Leu Thr
                       135
Asp Ser Glu Phe Lys Asp Pro Glu Thr Gly Lys Thr Cys Leu Leu Lys
                   150
                                        155
Ala Met Leu Asn Leu His Asn Gly Gln Asn Asp Thr Ile Ala Leu Leu
               165
                                   170
Leu Asp Val Ala Arg Lys Thr Asp Ser Leu Lys Gln Phe Val Asn Ala
                              185
Ser Tyr Thr Asp Ser Tyr Tyr Lys Gly Gln Thr Ala Leu His Ile Ala
                           200
                                                205
Ile Glu Arg Arg Asn Met Thr Leu Val Thr Leu Leu Val Glu Asn Gly
                        215
Ala Asp Val Gln Ala Ala Ala Asn Gly Asp Phe Phe Lys Lys Thr Lys
                    230
                                        235
Gly Arg Pro Gly Phe Tyr Phe Gly Glu Leu Pro Leu Ser Leu Ala Ala
                                    250
Cys Thr Asn Gln Leu Ala Ile Val Lys Phe Leu Leu Gln Asn Ser Trp
           260
                               265
                                                    270
Gln Pro Ala Asp Ile Ser Ala Arg Asp Ser Val Gly Asn Thr Val Leu
                            280
                                                285
His Ala Leu Val Glu Val Ala Asp Asn Thr Val Asp Asn Thr Lys Phe
                        295
                                            300
Val Thr Ser Met Tyr Asn Glu Ile Leu Ile Leu Gly Ala Lys Leu His
                                       315
                   310
Pro Thr Leu Lys Leu Glu Glu Ile Thr Asn Arg Lys Gly Leu Thr Pro
               325
                                   330
Leu Ala Leu Ala Ala Ser Ser Gly Lys Ile Gly Val Leu Ala Tyr Ile
           340
                                345
Leu Gln Arg Glu Ile His Glu Pro Glu Cys Arg His Leu Ser Arg Lys
                            360
Phe Thr Glu Trp Ala Tyr Gly Pro Val His Ser Ser Leu Tyr Asp Leu
                        375
                                            380
Ser Cys Ile Asp Thr Cys Glu Lys Asn Ser Val Leu Glu Val Ile Ala
                   390
                                        395
Tyr Ser Ser Ser Glu Thr Pro Asn Arg His Asp Met Leu Leu Val Glu
                                    410
Pro Leu Asn Arg Leu Leu Gln Asp Lys Trp Asp Arg Phe Val Lys Arg
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425
           420
Ile Phe Tyr Phe Asn Phe Phe Val Tyr Cys Leu Tyr Met Ile Ile Phe
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Thr Ala Ala Ala Tyr Tyr Arg Pro Val Glu Gly Leu Pro Pro Tyr Lys
                      455
                                           460
Leu Lys Asn Thr Val Gly Asp Tyr Phe Arg Val Thr Gly Glu Ile Leu
                                      475
                  470
Ser Val Ser Gly Gly Val Tyr Phe Phe Phe Arg Gly Ile Gln Tyr Phe
               485
                                   490
                                                      495
Leu Gln Arg Arg Pro Ser Leu Lys Ser Leu Phe Val Asp Ser Tyr Ser
                              505
                                                  510
           500
Glu Ile Leu Phe Phe Val Gln Ser Leu Phe Met Leu Val Ser Val Val
      515
                          520
Leu Tyr Phe Ser Gln Arg Lys Glu Tyr Val Ala Ser Met Val Phe Ser
                      535
                                          540
Leu Ala Met Gly Trp Thr Asn Met Leu Tyr Tyr Thr Arg Gly Phe Gln
                                     555
                  550
Gln Met Gly Ile Tyr Ala Val Met Ile Glu Lys Met Ile Leu Arg Asp
               565
                                   570
                                                       575
Leu Cys Arg Phe Met Phe Val Tyr Leu Val Phe Leu Phe Gly Phe Ser
                               585
                                                  590
           580
Thr Ala Val Val Thr Leu Ile Glu Asp Gly Lys Asn Asn Ser Leu Pro
                           600
Met Glu Ser Thr Pro His Lys Cys Arg Gly Ser Ala Cys Lys Pro Gly
                       615
                                          620
Asn Ser Tyr Asn Ser Leu Tyr Ser Thr Cys Leu Glu Leu Phe Lys Phe
                  630
                                      635
Thr Ile Gly Met Gly Asp Leu Glu Phe Thr Glu Asn Tyr Asp Phe Lys
                                   650
               645
Ala Val Phe Ile Ile Leu Leu Leu Ala Tyr Val Ile Leu Thr Tyr Ile
                                                  670
          660
                              665
Leu Leu Asn Met Leu Ile Ala Leu Met Gly Glu Thr Val Asn Lys
      675
                          680
Ile Ala Gln Glu Ser Lys Asn Ile Trp Lys Leu Gln Arg Ala Ile Thr
                      695
                               700
Ile Leu Asp Thr Glu Lys Ser Phe Leu Lys Cys Met Arg Lys Ala Phe
                                       715
                   710
Arg Ser Gly Lys Leu Gln Val Gly Phe Thr Pro Asp Gly Lys Asp
               725
                                   730
                                                       735
Asp Tyr Arg Trp Cys Phe Arg Val Asp Glu Val Asn Trp Thr Thr Trp
                               745
           740
Asn Thr Asn Val Gly Ile Ile Asn Glu Asp Pro Gly Asn Cys Glu Gly
                           760
                                              765
Val Lys Arg Thr Leu Ser Phe Ser Leu Arg Ser Gly Arg Val Ser Gly
                       775
                                          780
Arg Asn Trp Lys Asn Phe Ala Leu Val Pro Leu Leu Arg Asp Ala Ser
                   790
                                       795
Thr Arg Asp Arg His Ala Thr Gln Glu Glu Val Gln Leu Lys His
               805
                                 810
                                                   815
Tyr Thr Gly Ser Leu Lys Pro Glu Asp Ala Glu Val Phe Lys Asp Ser
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Met Val Pro Gly Glu Lys
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<213> Artificial Sequence

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